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## SEQUENCE LISTING

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## <120> GENE CLUSTER INVOLVED IN NOGALAMYCIN BIOSYNTHESIS, AND ITS USE IN PRODUCTION OF HYBRID ANTIBIOTICS

<130> 1574/49849

<150> PCT/FI99/00870

<151> 1999-10-20

<160> 18

<170> PatentIn version 3.0

<210>1

<211> 16020

<212> DNA

<213> Streptomyces nogalater ATCC 27451

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<221> misc feature

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<223> "overlapping sequence in the genes snoaM and snogN"

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<222> 6334..6356

<223> "overlapping sequence in the genes snoaG and snogC"

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tggaggtgct gctcaaggag aacaaggaga aggacgcctc ggtccccacc gccccgcacc 15840
acgatgcgtt cgccttcccg ttctccaccg ccggcaccgc cctgacggcg tgggtcgcgc 15900
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<210> 2

<211>342

<212> PRT

<213> Streptomyces nogalater ATCC 27451

<220>

<223> "translate of snogI, function: aminotransferase"

<400> 2

Met Thr Val His Val Trp Asp Tyr Leu Pro Glu Tyr Glu Leu Glu Arg
1 5 10 15

Glu Asp Ile His Asp Ala Val Glu Thr Val Phe Arg Ser Gly Arg Leu 20 25 30

Val Leu Gly Glu Ser Val Arg Gly Phe Glu Ser Glu Phe Ala Ser Phe 35 40 45

Gln Gly Val Gly His Ala Val Gly Val Asp Asn Gly Thr Asn Ala Val 50 55 60

Lys Leu Gly Leu Gln Ala Leu Gly Val Gly Pro Gly Asp Glu Val Val 65 70 75 80

Thr Val Ser Asn Thr Ala Ala Pro Thr Val Val Ala Ile Asp Ser Ala 85 90 95

Gly Ala Thr Pro Val Phe Val Asp Val Arg Glu Glu Asp Tyr Leu Met 100 105 110

Asp Thr Ser Gln Val Glu Ala Val Leu Thr Pro Arg Thr Arg Cys Leu 115 120 125

Leu Pro Val His Leu Tyr Gly Gln Cys Val Asp Met Ala Pro Leu Arg 130 135 140

Asp Leu Ala Ala Arg His Asn Leu Val Ile Leu Glu Asp Cys Ala Gln

155

160

Ala His Gly Ala Arg Arg His Gly Arg Leu Ala Gly Ser Thr Gly Asp 165 170 175

Ala Ala Ala Phe Ser Phe Tyr Pro Thr Lys Val Leu Gly Ala Tyr Gly
180 185 190

Asp Gly Gly Ala Val Leu Thr Asp Asp Glu Arg Val Ala Asp Arg Leu 195 200 205

Arg Arg Leu Arg Tyr Tyr Gly Met Glu Ser Arg Tyr Tyr Val Val Glu 210 215 220

Thr Pro Gly His Asn Ser Arg Leu Asp Glu Val Gln Ala Glu Ile Leu 225 230 235 240

Arg Arg Lys Leu Ser Arg Leu Pro Ser Tyr Ile Glu Ala Arg Arg Ala 245 250 255

Val Ala Arg Arg Tyr Glu Glu Gly Leu Ala Asp Thr Gly Leu Leu Leu 260 265 270

Pro Arg Thr Ala Gln Gly Asn Glu His Val Tyr Tyr Val Tyr Val Val 275 280 285

Arg His Pro Arg Arg Asp Ala Val Leu Glu Ala Leu Arg Ala Ser Tyr 290 295 300

Asp Ile Ala Leu Asn Ile Ser Tyr Pro Trp Pro Val His Thr Met Thr 305 310 315 320

Gly Phe Ser His Leu Gly Tyr Ala Lys Gly Ser Leu Pro Val Thr Glu 325 330 335

Ala Leu Ala Asp Glu Ile 340

<210>3

<211> 293

<212> PRT

<213> Streptomyces nogalater ATCC 27451

<220>

<223> "translate of snogJ, function: dTDP-glucose synthase"

<400> 3

Val Lys Gly Ile Ile Leu Ala Gly Gly Thr Gly Ser Arg Leu His Pro 1 5 10 15
Thr Thr Leu Ala Val Ser Lys Gln Leu Leu Pro Val Gly Asp Lys Pro 20 25 30
Met Ile Tyr Tyr Pro Leu Ser Val Leu Met Leu Ala Gly Val Thr Asp 35 40 45
Ile Leu Ile Ile Ser Thr Pro His Glu Leu Pro Arg Met Arg Arg Leu 50 55 60
Phe Gly Asp Gly Ala Gln Leu Gly Leu Arg Leu Ala Tyr Ala Glu Gln 65 70 75 80
Glu Lys Pro Arg Gly Ile Ala Glu Ala Phe Leu Ile Gly Ala Asp His 85 90 95
Val Gly Ser Asp Ala Val Ala Leu Ala Leu Gly Asp Asn Ile Phe His 100 105 110
Gly Ser Ser Phe Gln Gly Val Leu Arg Lys Glu Ala Glu Glu Leu Asp 115 120 125
Gly Cys Val Leu Phe Gly Tyr Pro Val Lys Asp Pro Gln Arg Tyr Gly 130 135 140
Val Gly Glu Ala Asn Ala Ser Gly Arg Leu Val Ser Ile Glu Glu Lys 145 150 155 160
Pro Val Arg Pro Arg Ser Asn Arg Ala Ile Thr Gly Leu Tyr Phe Tyr 165 170 175
Asp Asn Glu Val Val Asp Ile Ala Arg Arg Leu Arg Pro Ser Ala Arg 180 185 190
Gly Glu Leu Glu Ile Thr Asp Ile Asn Arg Thr Tyr Met Glu Arg Gly 195 200 205
Arg Ala Arg Leu Val Asp Leu Gly Arg Gly Phe Ala Trp Leu Asp Thr 210 215 220
Gly Thr Pro Glu Ser Leu Leu Gln Ala Ser Gln Tyr Val Ser Ala Leu 225 230 235 240
Glu Glu Arg Gln Gly Ile Arg Ile Ala Cys Ile Glu Glu Val Ala Leu

Arg Met Gly Phe Ile Asn Ala Gln Ala Cys Tyr Glu Leu Gly Ala Arg 260 265 270

Leu Ser Gly Ser Gly Tyr Gly Gln Tyr Val Met Ala Ile Ala Glu Glu 275 280 285

Cys Thr Gly Arg Val 290

<210>4

<211>238

<212> PRT

<213> Streptomyces nogalater ATCC 27451

<220>

<223> "translate of snogA, function: aminomethyl transferase"

<400>4

Val Tyr Gly Arg Glu Leu Ala Asp Val Tyr Glu Met Val Tyr Arg Ser 1 5 10 15

Arg Gly Lys Ser Trp Ala Asp Glu Ala Glu Arg Val Thr Ala Glu Ile 20 25 30

Arg Ser Arg Arg Pro Gly Ala Arg Ser Leu Leu Asp Val Ala Cys Gly 35 40 45

Thr Gly Ala His Leu Glu Ala Phe Arg Gly Leu Phe Ala His Thr Glu 50 55 60

Gly Leu Glu Leu Ser Asp Glu Met Arg Ala Leu Ala Glu Arg Arg Leu 65 70 75 80

Pro Gly Val Pro Val Arg Pro Gly Asp Met Arg Asp Phe Ala Leu Ser 85 90 95

Gly Arg Phe Asp Ala Val Val Cys Leu Phe Cys Ser Ile Gly Tyr Leu
100 105 110

Glu Thr Val Ala Asp Met Arg Ala Ala Val Arg Thr Met Ala Ala His
115 120 125

Leu Val Pro Gly Gly Val Leu Val Val Glu Pro Trp Phe Pro Glu 130 135 140

Arg Phe Leu Glu Gly Tyr Val Ala Gly Asp Leu Ala Arg Gly Glu Gly 145 150 155 160
Arg Thr Val Ala Arg Val Ser His Ser Thr Arg Gln Gly Arg Arg Thr 165 170 175
Arg Met Glu Val Arg Phe Leu Val Gly Glu Ala Thr Gly Ile Arg Glu 180 185 190
Phe Thr Glu Ile Asp Leu Leu Thr Leu Phe Thr Arg Glu Glu Tyr Leu 195 200 205
Ala Ala Phe Glu Asp Ala Gly Cys Pro Ala Glu Phe Leu Asp Asp Gly 210 215 220
Leu Thr Gly Arg Gly Leu Phe Val Gly Val Arg Gly Ala Gly 225 230 235
<210> 5 <211> 324 <212> PRT <213> Streptomyces nogalater ATCC 27451
<220> <223> "translate of snoaM, function: polyketide cyclase"
<400> 5
Met Thr Ala Ala Trp Gly Ala Pro Leu Tyr Pro Pro Trp Ile Pro Ala 1 5 10 15
Arg Pro Gly Arg Arg Cys Gly Ala Gly Arg Arg Val Arg Cys Pro 20 25 30
Pro Val Glu Pro Ala Ser Arg Pro Arg Gln Glu Gly Arg Val Ser Val 35 40 45
35 40 45  Val Pro Ala Leu Arg Gln Pro Ser Pro Ser Thr Asn Pro Glu Val Arg

His Met Cys Ala Glu Met Arg Glu His Phe Gly Val Glu Phe Ser Pro 100 105 110
Asp Glu Leu Pro Asp Gly Glu Phe Leu Ser Leu Asp Arg Ile Thr Leu 115 120 125
Thr Thr His Thr Gly Thr His Val Asp Ala Pro Ser His Tyr Gly Ser 130 135 140
Arg Ala Leu Tyr Gly Asp Gly Val Pro Arg His Ile Asp Gln Met Pro 145 150 155 160
Leu Glu Trp Phe Phe Gly Arg Gly Val Val Leu Asp Leu Thr Asp Ala 165 170 175
Pro Thr Gly Thr Val Ser Ala Ala Arg Leu Glu Lys Glu Leu Ala Arg 180 185 190
Thr Gly Cys Ala Leu Arg Pro Gly Asp Ile Val Leu Leu His Thr Gly 195 200 205
Ala Gln Arg His Ala Gly Thr Pro Arg Tyr Phe Thr Asp Phe Ala Gly 210 215 220
Leu Asp Gly Pro Ala Val Arg Met Leu Leu Asp His Gly Val Arg Val 225 230 235 240
Ile Gly Thr Asp Ala Phe Ser Leu Asp Ala Pro Phe Gly His Ile Ile 245 250 255
Asp Arg Tyr Arg Ala Thr Gly Asp Arg Ser Val Leu Trp Pro Ala His 260 265 270
Val Val Gly Arg Glu Arg Glu Tyr Cys Gln Ile Glu Arg Leu Ala Asn 275 280 285
Leu Asp Arg Leu Pro Val Ser Phe Gly Phe Arg Val Cys Cys Phe Pro 290 295 300
Val Lys Val Ala Gly Ala Gly Ala Gly Trp Thr Arg Ala Val Ala Leu 305 310 315 320
Val Asp Glu Asp
<210> 6 <211> 408

<212> PRT

<213> Streptomyces nogalater ATCC 27451 <220> <223> "translate of snogN, function: unknown" <400> 6 Met Val Met Lys Leu Thr Asp Ser Glu Leu Gly Arg Ala Leu Leu Ser Leu Arg Gly Tyr Gln Trp Leu Arg Gly Ile His His Asp Pro Tyr Ala 25 Leu Leu Arg Ala Glu Ser Asp Asp Pro Ala Gln Leu Gly Arg Leu 45 Leu Arg Glu Arg Gly Arg Leu His Arg Ser Asp Thr Gly Thr Trp Val 50 55 Thr Ala Asp His Ala Thr Ala Ser Arg Leu Leu Ala Asp Pro Arg Phe 70 75 80 65 Val Leu Arg Arg Pro Pro Ala Gly Pro Ala Thr Gly Thr Gly Asp Val 85 90 95 Met Pro Trp Glu Glu Ala Thr Leu Ser Asp Leu Leu Pro Leu Asp Glu 105 100 110 Ala Arg Leu Thr Thr Asp Arg Ala Arg Cys Arg Arg Leu Gly Ala Thr 115 120 125 Ala Ala Arg Ile Ala Ala Asp Gly Pro Val Ala Thr Arg Leu Ala Asp 135 140 Leu Ala Gly Ala Arg Ala Glu Gln Val Arg Ser Thr Gly His Phe Asp 150 155 Leu Arg Ala Asp Tyr Ala Leu Pro Tyr Ala Val Glu Pro Ala Cys Ala 175 170 Leu Leu Gly Leu Pro Ala Gly Gln Cys Ser Leu Phe Gly Ala Phe Ser 185 190 Pro Ala Val Leu Leu Asp Ala Thr Val Val Pro Pro Arg Leu Pro Glu 200 195 205

Ala Arg Ala Leu Ile Ala Ser Thr Ala Glu Leu Thr Ala Leu Trp Pro

Arg Leu Ala Pro Ser Leu Ser Lys Thr Val Pro Glu Asp Glu Ala Pro 

Asp Leu Phe Leu Leu Thr Ala Val Leu Leu Val Pro Ala Val Val His 

Leu Val Cys Glu Ala Val Ala Ala Leu Ser His Asp Pro Gly Gln Ala 

Gly Leu Aeu Arg Asp Asp Pro Val Leu Ala Ala Pro Ala Val Glu Glu 

Thr Leu Arg His Ala Pro Pro Ala Arg Leu Phe Thr Leu His Ala Thr 

Gly Pro Glu Arg Val Ala Asp Val Asp Leu Pro Ala Gly Ala Glu Val 

Ala Val Val Ala Ala Ala His Arg Asp Pro Ser Trp Cys Pro Asp 

Pro Asp Arg Phe Asp Leu Thr Arg Asn Glu Arg His Leu Ala Leu Pro 

Pro Asp Leu Pro Leu Gly Ala Leu Ala Pro Leu Leu Arg Val Cys Ala 

Thr Ala Ala Val Ala Ala Leu Ala Ala Gly Leu Leu Pro Leu Arg Ala 

Val Gly Pro Pro Val Arg Arg Leu Arg Ala Pro Val Thr Arg Ser Val 

Leu Arg Phe Pro Val Ala Pro Cys 

<210> 7

<211>422

<212> PRT

<213> Streptomyces nogalater ATCC 27451

<220>

<223> "translate of snoaG, function: hydroxylase"

<400> 7

Met Asp Asn Arg Glu Thr Val Arg Pro Val Ser Val Cys Arg Val Cys  1 5 15 15
Gly Gly Asn Asp Trp Gln Asp Val Val Asp Phe Gly Asp Val Pro Leu 20 25 30
Ala Asn Gly Phe Leu Ser Pro Ala Asp Ser Tyr Glu Asn Glu Arg Arg 35 40 45
Tyr Pro Leu Gly Val Leu Ser Cys Arg Ala Cys Arg Leu Met Ser Leu 50 55 60
Thr His Val Val Asp Pro Glu Val Leu Tyr Arg Asp Tyr Ala Tyr Thr 65 70 75 80
Thr Pro Asp Ser Glu Met Ile Thr Gln His Met Arg His Ile Thr Ala 85 90 95
Leu Cys Arg Thr Arg Phe Glu Leu Pro Pro Asp Ser Leu Val Val Glu 100 105 110
Leu Gly Ser Asn Thr Gly Arg Gln Leu Met Ala Phe Arg Glu Ala Gly 115 120 125
Met Arg Thr Leu Gly Val Asp Pro Ala Arg Asn Leu Thr Asp Val Ala 130 135 140
Arg Arg Asn Gly Ile Glu Thr Phe Pro Asp Phe Phe Ser His Asp Val 145 150 155 160
Ala Arg Thr Ile Arg Arg Asp His Gly Gln Ala Arg Leu Val Leu Gly 165 170 175
Arg His Val Phe Ala His Ile Asp Asp Val Ser Asp Ile Ala Ala Gly 180 185 190
Val Arg Glu Leu Leu Ser Pro Asp Gly Val Phe Ala Ile Glu Val Pro 195 200 205
Tyr Val Leu Asp Leu Leu Glu Lys Val Ala Phe Asp Thr Ile Tyr His 210 215 220
Glu His Leu Ser Tyr Phe Thr Met Arg Ser Phe Val Thr Leu Phe Ala 225 230 235 240
Arg His Gly Leu Arg Val Leu Asp Val Glu Arg Phe Gly Val His Gly



245 250 255

Gly Ser Val Leu Val Phe Val Gly His Glu Asp Gly Pro Trp Pro Glu 260 265 270

Arg Pro Ser Val Pro Glu Leu Leu Arg Val Glu Arg Gln Arg Gly Leu 275 280 285

Tyr Asp Asp Ala Thr Tyr Arg Thr Phe Ala Gln Arg Ile Glu Arg Val 290 295 300

Arg Thr Glu Leu Pro Glu Leu Leu Arg Ser Leu Val Ala Gln Gly Lys 305 310 315 320

Arg Ile Val Gly Tyr Gly Ala Pro Ala Lys Gly Asn Thr Ile Leu Thr 325 330 335

Val Cys Gly Leu Gly Leu Lys Glu Leu Glu Tyr Cys Thr Asp Thr Thr 340 345 350

Glu Leu Lys Gln Gly Arg Val Leu Pro Gly Thr His Ile Pro Val His 355 360 365

Ala Pro Glu His Ala Lys Glu His Ile Pro Asp Tyr Tyr Leu Leu Leu 370 375 380

Ala Trp Asn Tyr Ala Thr Glu Ile Leu Asp Lys Glu Thr Ala Phe Arg 385 390 395 400

Asp Asn Gly Gly Arg Phe Ile Val Pro Ile Pro Arg Pro Ser Ile Leu 405 410 415

Thr Ser Pro Ser Gly Ser 420

<210>8

<211> 291

<212> PRT

<213> Streptomyces nogalater ATCC 27451

<220>

<223> "translate of snogC, function: dTDP-4-dehydrorhamnose reductase"

<400> 8

Met Leu Ala Arg His Leu Thr Ala Ala Leu Ala Glu Thr Gly Arg Ser 1 5 10 15

Arg Pro Ala A 20	la Glu Ala Va 25	al Val Leu ( 30	Gly Arg Arg A	la Leu Asp Ile
Thr Asp Gly A	rg Ala Val A 40	sp Ala Ala 45	Phe Ala Ala H	is Arg Pro Arg
Val Val Val As 50	sn Cys Ala Al 55	la Phe Thr A	Asp Val Asp C	ily Ala Glu Ser
• .	lu Ala Met A 70	rg Val Asn 75	Gly Gly Gly F 80	ro Arg Leu Le
Ala Arg Arg C 85	ys Ala Arg H 90	-	Arg Leu Ile Hi 95	s Val Ser Thr
Asp Tyr Val Pl 100	he Pro Gly A 105		Ser Pro Tyr G 10	dy Glu Ser Asp
Ala Pro Gly Pr 115	o Arg Thr Va 120	al Tyr Gly A 125	Arg Ser Lys Le	eu Ala Gly Glu
Arg Ala Val Le 130	eu Ser Leu Lo 135	eu Pro Asp 140	Thr Gly Thr V	al Val Arg Thr
•	yr Gly Gly G 150	In Gly Arg	Ser Phe Val A 160	rg Thr Met Leu
Glu Arg Ala Pr 165	ro Asp Asp C	•	Asp Val Val A 175	sn Asp Gln Trp
Gly Gln Pro Tl 180	ır Trp Ala Gl 185	-	Ala Arg Leu L 90	eu Val Thr Leu
Ala Arg Thr Pr 195	o Pro Asp A 200	rg Ala Arg 205	Gly Ile Phe Hi	s Ala Thr Asn
Ala Gly Ala Al 210	a Thr Trp Ty 215	r Glu Leu A 220	Ala Arg Glu V	al Phe Arg Leu
	sp Pro Glu Ai 230	rg Val Arg	Pro Val Ala T 240	hr Ala Asp Arg

Pro Gly Pro Ala Pro Arg Pro Ala Cys Thr Val Leu Gly His Asp Arg 245 250 255

Trp Arg Leu Val Gly Val Ala Pro Pro Arg Asp Trp Arg Ala Ala Leu

260 265

Arg Glu Ala Met Arg Gln Leu Leu Pro Gly Gly Arg Leu Arg Asn Leu 275 280 285

270

Thr Gly Thr 290

<210> 9

<211>350

<212> PRT

<213> Streptomyces nogalater ATCC 27451

<220>

<223> "translate of snogK, function: dTDP-glucose-4,6-dehydratase"

<400> 9

Met Ala Ser His Thr Ser Ala Thr Thr Asp Val Asn Ile Leu Val Thr 1 5 10 15

Gly Ala Val Gly Phe Ile Gly Ser Ala Tyr Val Arg Met Leu Leu Glu 20 25 30

Asn Arg Ala Pro Gly Ala Gly Ala Pro Ala Val Arg Val Thr Val Leu 35 40 45

Asp Lys Leu Thr Tyr Ala Gly Asn Leu Thr Asn Leu Asp Ala Val Arg 50 55 60

Gly Asp Arg Leu Arg Phe Val Arg Gly Asp Ile Leu Asp Ala Glu Leu 65 70 75 80

Val Asp Glu Leu Met Ala His Ser Asp Gln Val Val His Phe Ala Ala 85 90 95

Glu Ser His Val Asp Arg Ser Ile Arg Ala Ala Asp Asp Phe Val Leu 100 105 110

Thr Asn Val Val Gly Thr Gln Arg Leu Leu Asp Ala Ala Leu Arg His
115 120 125

Gly Val Glu Pro Phe Val Leu Val Ser Thr Asp Glu Val Tyr Gly Ser 130 135 140

Ile Ala Ser Gly Ser Trp Pro Glu Glu His Pro Leu Ser Pro Asn Ser 145 150 155 160

Pro Tyr Ala Ala Ser Lys Ala Ser Ala Asp Leu Met Ala Phe Ala Cys 165 170 175
His Arg Thr His Gly Leu Asp Val Arg Val Thr Arg Cys Ser Asn Asn 180 185 190
Tyr Gly Pro Arg Gln His Pro Glu Lys Leu Ile Pro Arg Phe Val Thr 195 200 205
Asn Leu Leu Asp Gly Leu Pro Val Pro Leu Tyr Gly Asp Gly Arg As 210 . 215 220
Val Arg Glu Trp Leu His Val Glu Asp His Cys Arg Gly Val Asp Leu 225 230 235 240
Val Arg Thr Ala Gly Arg Pro Gly Gly Val Tyr His Ile Gly Gly Gly 245 250 255
Arg Glu Leu Ser Asn Arg Glu Leu Val Gly Met Leu Leu Glu Leu Cy 260 265 270
Gly Ala Asp Trp Ser Ser Val Arg His Val Pro Asp Arg Lys Gly His 275 280 285
Asp Leu Arg Tyr Ser Leu Asp Trp Gly Arg Ala Arg Glu Glu Leu Gly 290 295 300
Tyr Arg Pro Ala Arg Glu Phe Ser Ser Gly Leu Arg Ser Thr Val Gln 305 310 315 320
Trp Tyr Arg Glu Asn Arg Ser Trp Trp Glu Pro Leu Lys Arg Gly Val 325 330 335
Thr Ala Pro Gly Gly Thr Ser Thr Val Val Pro Gly Val Arg 340 345 350
<210> 10 <211> 134 <212> PRT <213> Streptomyces nogalater ATCC 27451
<220> <223> "translate of snoaL, function: NAME cyclase"
<400> 10

_	er Ala Phe A 5	asn Thr Gly 10	Arg Thr Asp Asp 15	Val Asp Glu Tyr
Ile His Pro 20		u Asn Pro 2 25	Ala Thr Leu Glu F	lis Gly Ile His
Thr Gly Pro	o Lys Ala Pl 40		Leu Val Gly Trp V 45	al Arg Ala Thr
Phe Ser Glo	u Glu Ala A 55	rg Leu Glu 60	Glu Val Arg Ile G	ilu Glu Arg Gly
Pro Trp Va 65	ll Lys Ala T 70	yr Leu Val	Leu Tyr Gly Arg I 80	His Val Gly Arg
	y Met Pro P 35	ro Thr Asp 90	Arg Arg Phe Ser 95	Gly Glu Gln Val
His Leu Mo	•	al Asp Gly	Lys Ile Arg Asp H 110	is Arg Asp Trp
Pro Asp Ph	ie Gln Gly T 12	_	Gln Leu Gly Asp 125	Pro Trp Pro Asp
Asp Glu Gl 130	y Trp Arg F	Pro		
<210> 11 <211> 235 <212> PR7 <213> Stre	r ptomyces n	ogalater Al	CCC 27451	
<220> <223> "tra	nslate of sno	oK, function	n: unknown"	
<400> 11				
-	sp Pro Gly C 5	Gly Pro Thr 10	Thr Ala Glu Asn	Leu Ser Lys Glu
Ala Val Ar 20		arg Glu Gln 25	Gly Tyr Val His I 30	le Pro Arg Val

Leu Glu Lys Glu Gly Arg Glu Ile Ser Gly Ile Ala Leu Arg Leu Ala

Leu Ser Glu Thr Glu Val Thr Ala Phe Arg Ala Ala Cys Glu Glu Val $35 \hspace{1cm} 40 \hspace{1cm} 45$ 

50	55	60			
Gly Ala Pro I 65	eu Arg Val	Tyr Ser Se 75	r Asp Ile Lei 80	ı Val Lys Glu Pro	
Lys Arg Thr 1		· Leu Val H 90	is Asp Asp ( 95	Glu Thr Gly Leu Pro	Э
Leu Asn Glu 100	Leu Ser Ala 10		hr Ala Trp Il 110	e Ala Leu Thr Asp	
Val Pro Val C 115	Glu Arg Gly 120	-	er Tyr Val Pi 25	ro Gly Ser His Leu	
Arg Ala Arg	Glu Asp Arg 135	g Gln Glu H 140	lis Met Thr S	Ser Phe Ala Glu Pho	Э
Arg Asp Leu 145	Ala Asp Va 150	l Trp Pro A 155	asp Tyr Pro 1 160	Ггр Gln Pro Arg Va	ıl
Ala Val Pro V	_	Gly Asp Va 170	al Val Phe Hi 175	is His Cys Arg Thr	
Val His Met A	Ala Glu Ala 18		er Asp Ser V 190	al Arg Met Ala His	
Gly Val Val 7 195	Tyr Met Asp 200	-	la Thr Tyr A 05	rg Pro Gly Val Gln	•
Asp Gly His I 210	Ceu Ser Arg 215	Leu Ser Pr 220	o Gly Asp P	ro Leu Glu Gly Glu	l
Leu Phe Pro 1 225	Leu Val Thr 230	Ala Gly Tl 235	nr Arg Gln		
<210> 12 <211> 390 <212> PRT <213> Strept	omyces nog	alater ATC	C 27451		
<220> <223> "transl	late of snogl	D, function:	glycosyl tra	nsferase"	
<400> 12					

Met Arg Val Pro Gly Ser Cys Arg Thr Gly Gly Ile Met Arg Ala Leu 1 5 10 15

Page 27

Phe Ile Thr Ser Pro Gly Leu Ser His Ile Leu Pro Thr Val Pro Leu 20 25 30
Ala Gln Ala Leu Arg Ala Leu Gly His Glu Val Arg Tyr Ala Thr Gly 35 40 45
Gly Asp Ile Arg Ala Val Ala Glu Ala Gly Leu Cys Ala Val Asp Val 50 55 60
Ser Pro Gly Val Asn Tyr Ala Lys Leu Phe Val Pro Asp Asp Thr Asp 65 70 75 80
Val Thr Asp Pro Met His Ser Glu Gly Leu Gly Glu Gly Phe Phe Ala 85 90 95
Glu Met Phe Ala Arg Val Ser Ala Val Ala Val Asp Gly Ala Leu Arg 100 105 110
Thr Ala Arg Ser Trp Arg Pro Asp Leu Val Val His Thr Pro Thr Gln 115 120 125
Gly Ala Gly Pro Leu Thr Ala Ala Ala Leu Gln Leu Pro Cys Val Glu 130 135 140
Leu Pro Leu Gly Pro Ala Asp Ser Glu Pro Gly Leu Gly Ala Leu Ile 145 150 155 160
Arg Arg Ala Met Ser Lys Asp Tyr Glu Arg His Gly Val Thr Gly Glu 165 170 175
Pro Thr Gly Ser Val Arg Leu Thr Thr Thr Pro Pro Ser Val Glu Ala 180 185 190
Leu Leu Pro Glu Asp Arg Arg Ser Pro Gly Ala Trp Pro Met Arg Typ 195 200 205
Val Pro Tyr Asn Gly Gly Ala Val Leu Pro Asp Trp Leu Pro Pro Ala 210 215 220
Ala Gly Arg Arg Ile Ala Val Thr Leu Gly Ser Ile Asp Ala Leu 225 230 235 240

Val Asp Ala Glu Phe Val Leu Thr Leu Gly Gly Gly Asp Leu Ala Leu

Ser Gly Gly Ile Ala Lys Leu Ala Pro Leu Phe Ser Glu Val Ala Asp

250

245

255

Leu Val Ala Leu Ala Gly 385 390

<210> 13

<211> 275

<212> PRT

<213> Streptomyces nogalater ATCC 27451

<220>

<223> "translate of snoW, function: unknown"

<400> 13

Met Thr Val Leu Val Thr Gly Ala Thr Gly Asn Val Gly Arg His Val 1 5 10 15

Val Thr Gly Leu Leu Ala Ala Gly Arg Arg Val Arg Ala Leu Thr Arg 20 25 30

Thr Pro Asp Arg Ser Gly Leu Pro Gly Gly Ala Glu Ile Thr Gly Gly 35 40 45

Asp Leu Thr Arg Pro Glu Thr Tyr Glu Arg Met Leu Asp Gly Val Glu 50 55 60

Ala Val Tyr Leu Phe Pro Val Pro Glu Thr Ala Ala Ala Phe Ala Gly Ala Ala Arg Arg Ala Gly Val Arg Arg Ile Val Val Leu Ser Ser Asp Ser Val Thr Asp Gly Thr Asp Thr Gly Gly His Arg Arg Val Glu Leu Ala Val Glu Asp Thr Gly Leu Glu Trp Thr His Val Arg Pro Gly Glu Phe Ala Leu Asn Lys Val Thr Leu Trp Ala Pro Ser Ile Arg Ala Glu Gly Val Val Arg Ser Ala Tyr Pro Asp Ala Arg Val Ala Pro Val His Glu Ala Asp Val Ala Ala Val Ala Val Thr Ala Leu Leu Lys Glu Gly His Ala Gly Arg Ala Tyr Ser Val Thr Gly Pro Gln Ala Leu Thr Gln Arg Glu Gln Val Arg Ala Val Gly Glu Gly Leu Gly Arg Ser Leu Ala Phe Val Glu Val Thr Pro Gly Gln Ala Arg Ala Asp Leu Thr Ala Gln Gly Leu Pro Ala Pro Ile Ala Asp Tyr Val Leu Ala Phe Gln Ala Gly Trp Thr Glu Arg Pro Ala Pro Ala Arg Pro Thr Val Arg Glu Val Thr Gly Arg Pro Ala Arg Thr Leu Ala Gln Trp Ala Ala Asp His Arg Ala Asp Phe Arg <210> 14 <211>424 <212> PRT <213> Streptomyces nogalater ATCC 27451

<220> <223> "translate of snogE, function: glycosyl transferase"
<400> 14
Val Arg Val Leu Leu Thr Ser Phe Ala Met Asp Ala His Phe Cys Thr 1 5 10 15
Ala Val Pro Leu Ala Trp Ala Leu Arg Ser Ala Gly His Glu Val Arg 20 25 30
Val Ala Gly Gln Pro Ala Leu Thr Ser Thr Ile Thr Gly Ala Gly Leu 35 40 45
Thr Ala Val Pro Val Gly Arg Asp His Thr His Gly Ser Leu Leu Gly 50 55 60
Arg Val Gly Ser Asp Ile Leu Ala Leu His Asp Glu Ala Asp Tyr Leu 65 70 75 80
Glu Ala Arg His Asp Ala Leu Gly Phe Glu Phe Leu Lys Gly His Asn 85 90 95
Thr Val Met Ser Ala Leu Phe Tyr Ser Gln Ile Asn Asn Asp Ser Met 100 105 110
Val Asp Asp Leu Val Asp Phe Ala Arg His Trp Arg Pro Asp Leu Val 115 120 125
Val Trp Glu Pro Phe Thr Phe Ala Gly Ala Val Ala Ala Arg Ala Ser 130 135 140
Gly Ala Ala His Ala Arg Leu Leu Ser Phe Pro Asp Leu Phe Leu Ser 145 150 155 160
Thr Arg Arg Leu Phe Leu Glu Arg Met Ala Arg Gln Glu Pro Glu His

Thr Arg Arg Leu Phe Leu Glu Arg Met Ala Arg Gln Glu Pro Glu His 165 170 175

His Asp Asp Thr Leu Ala Glu Trp Leu Asp Trp Thr Leu Gly Arg His 180 185 190

Gly His Ser Phe Asp Glu Glu Ile Val Thr Gly Gln Trp Ser Ile Asp 195 200 205

Gln Thr Pro Ala Pro Val Arg Leu Asp Ala Gly Gly Pro Thr Val Pro 210 215 220

Met Arg Tyr Val Pro Tyr Ser Gly Leu Val Pro Thr Val Val Pro Asp 225 230 235 240
Trp Leu Arg Arg Pro Pro Glu Arg Pro Arg Val Leu Val Thr Leu Gly 245 250 255
Ile Thr Ser Arg Arg Val Lys Ser Phe Leu Ala Val Ser Val Asp Asp 260 265 270
Leu Phe Glu Ala Val Ala Gly Leu Gly Val Glu Val Val Ala Thr Leu 275 280 285
Asp Ala Asp Gln Arg Glu Leu Leu Gly Arg Val Pro Asp His Phe Arg 290 295 300
Ile Val Glu His Val Pro Leu Asp Ala Val Leu Pro Thr Cys Ser Ala 305 310 315 320
Ile Val His His Gly Gly Ala Gly Thr Trp Ser Thr Ala Ala Val Tyr 325 330 335
Gly Val Pro Gln Val Ser Leu Gly Ser Met Trp Asp His Phe Tyr Arg 340 345 350
Ala Arg Arg Leu Glu Glu Leu Gly Ala Gly Leu Arg Leu Pro Ser Gly 355 360 365
Glu Leu Thr Ala Glu Gly Leu Arg Thr Arg Leu Glu Arg Val Leu Gly 370 375 380
Glu Pro Ser Phe Gly Thr Ala Ala Gln Ala Leu Ser Asp Thr Ile Ala 385 390 395 400
Ala Glu Pro Ser Pro Ser Glu Val Val Pro Val Leu Glu Glu Leu Thr 405 410 415
Gly Arg His Arg Pro Gly Thr Arg 420
<210> 15 <211> 139 <212> PRT <213> Streptomyces nogalater ATCC 27451
<220> <223> "translate of snoL, function: unknown"

1	5
	1

Met Ser Thr Thr Ala Asn Lys Glu Arg Cys Leu Glu Met Val Ala Ala 1 5 10 15

Trp Asn Arg Trp Asp Val Ser Gly Val Val Ala His Trp Ala Pro Asp 20 25 30

Val Val His Tyr Asp Asp Glu Asp Lys Pro Val Ser Ala Glu Glu Val 35 40 45

Val Arg Arg Met Asn Ser Ala Val Glu Ala Phe Pro Asp Leu Arg Leu 50 55 60

Asp Val Arg Ser Ile Val Gly Glu Gly Asp Arg Val Met Leu Arg Ile 65 70 75 80

Thr Cys Ser Ala Thr His Gln Gly Val Phe Met Gly Ile Ala Pro Thr
85 90 95

Gly Arg Lys Val Arg Trp Thr Tyr Leu Glu Glu Leu Arg Phe Ser Glu
100 105 110

Ala Gly Lys Val Val Glu His Trp Asp Val Phe Asn Phe Ser Pro Leu 115 120 125

Phe Arg Asp Leu Gly Val Val Pro Asp Gly Leu 130 135

<210> 16

<211> 155

<212> PRT

<213> Streptomyces nogalater ATCC 27451

<220>

<223> "translate of snoO, function: homologous to mtmX of mithramycin cluster"

<400> 16

Met Ser Val Arg Thr Asp Gln Thr Ala Ala Pro Glu Asp Arg Ala Ala 1 5 10 15

Ala Thr Asp Pro Gly Phe Gly His Leu Tyr Ala Gln Val Gln Gln Phe 20 25 30

Tyr Ala Arg Gln 35	Met Gln Leu L 40	eu Asp Se 45	er Gly Ala Ala Glu Glu Tı	г <b>р</b>				
Ala Ala Thr Phe 50		y Thr Phe 0	e Ala Arg Pro Ser Ser Pro	)				
Glu Pro Ala Arg 65 70	•	ı Leu Ala	Ala Gly Ala Arg Ala Ala 80	l				
Ala Glu Arg Leu 85	Ala Ala Glu Gl 90	y Leu Ser 95	His Arg His Val Ile Gly					
Met Thr Ala Val 100	Arg Arg Glu Pi 105	ro Asp Gl	y Ser Val Phe Val Arg Se	er				
Tyr Ala Gln Val 115	Phe Ala Thr Arg 120	g Arg Gly 125	Glu Ala Pro Arg Leu Hi	S				
Leu Ile Cys Val ( 130	•	l Leu Val 140	Arg Glu Gly Pro Gly Let	u				
Lys Val Arg Glu 145 15	_	_	Ala					
<210> 17 <211> 281 <212> PRT <213> Streptomy	yces nogalater A	ATCC 274	<b>1</b> 51					
<220> <223> "translate of snoaF, function: C-7 ketoreductase"								
<400> 17								
Val Arg Ala Met 1 5	Thr Asp Ser Th	nr Gly Pro 15	Arg Pro Val Pro Ala Me	et				
Ser Pro Ala Pro 3 20	Ser Pro Thr Pro 25	Ser Pro 0	Gly Pro Ala Pro Gly Ser					
Glu Pro Ala Pro Leu Ala Val Ile Val Thr Gly Gly Gly Ser Gly Ile 35 40 45								

Val Val Gly Arg Thr Glu Asp Ala Leu Ala Gln Thr Ala Glu Gly Cys

Gly Arg Ala Thr Ala Arg Ala Phe Ala Ala Gln Gly Ala Lys Val Leu 50 55 60

65	70	75	80			
Ala Asp Met 85	_	_	Val Ala Ser Pro 95	Asp Gly Pro		
Gln Ala Val	Val Asn Ala A 105	-	Glu Phe Gly Arg 10	Ile Asp Val		
Leu Val Asn 115	Asn Ala Ala 120	Val Ala Gly 1 125	Met Glu Thr Leu	Gln Thr Val		
Asp Arg Asp 130	Ala Val Ala 135	Arg Gln Phe 140	Gly Thr Asn Leu	Thr Ala Pro		
Leu Phe Leu 145	Val Gln Ser . 150	Ala Leu Gly 155	Ala Leu Glu Lys 160	Ser Arg Gly		
Ile Val Val Asn Val Gly Thr Ala Ala Thr Leu Gly Leu Arg Ala Ala 165 170 175						
Pro Thr Gly Ala Leu Tyr Gly Ala Ser Lys Val Ala Leu Asp Tyr Leu 180 185 190						
Thr Arg Thr 195	Trp Ala Val ( 200	Glu Leu Ala 205	Pro Arg Gly Ile A	arg Val Val		
Gly Val Ala Pro Gly Val Ile Asp Thr Gly Ile Gly Val Arg Met Gly 210 215 220						
Met Thr Pro 225	Glu Gly Tyr 230	Arg Glu Phe 235	Leu Thr Gly Met 240	Gly Gly Arg		
Val Pro Val ( 24		Gly Arg Pro ( 250	Glu Asp Val Ala ' 255	Trp Trp Ile		
Val Gln Leu 260	Ala Arg Pro 0 265		Tyr Ala Thr Gly 1 70	Met Val Val		
Pro Val Asp Gly Gly Leu Ser Leu Val 275 280						

<210> 18

<211> 190

<212> PRT

<213> Streptomyces nogalater ATCC 27451

<220>

<223> "translate of snoN, function: unknown" <400> 18 Val Gln Glu Thr Glu Pro Gly Val Pro Ala Asp Leu Pro Ala Glu Ser Asp Pro Ala Ala Leu Glu Arg Leu Ala Ala Arg Tyr Arg Arg Asp Gly Tyr Val His Val Pro Gly Val Leu Asp Ala Gly Glu Val Ala Glu Tyr Leu Ala Glu Ala Arg Arg Leu Leu Ala His Glu Glu Ser Val Arg Trp Gly Ser Gly Ala Gly Thr Val Met Asp Tyr Val Ala Asp Ala Gln Leu Gly Ser Asp Thr Met Arg Arg Leu Ala Thr His Pro Arg Ile Ala Ala Leu Ala Glu Tyr Leu Ala Gly Ser Pro Leu Arg Leu Phe Lys Leu Glu Val Leu Leu Lys Glu Asn Lys Glu Lys Asp Ala Ser Val Pro Thr Ala Pro His His Asp Ala Phe Ala Phe Pro Phe Ser Thr Ala Gly Thr Ala Leu Thr Ala Trp Val Ala Leu Val Asp Val Pro Val Glu Arg Gly Cys Met Thr Phe Val Pro Gly Ser His Leu Leu Pro Asp Pro Asp Thr Gly 

Asp Glu Pro Trp Ala Gly Ala Phe Thr Arg Pro Gly Glu Ile